083,913 10/477,971

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(FILE 'CAPLUS' ENTERED AT 12:26:22 ON 23 JAN 2004) DEL HIS

FILE 'USPATFULL' ENTERED AT 12:38:05 ON 23 JAN 2004

L10 S PHARMACEUTICAL (3W) KIT#/IA

L2 3052 S PHARMACEUTICAL (3W) KIT#

L3 1 S STOLL, ANDREW/IN

L41 S US6344482/PN

FILE 'WPIDS' ENTERED AT 12:41:38 ON 23 JAN 2004

L51 S WO9739759/PN

ANSWER 1 OF 1 WPIDS COPYRIGHT 2004 THOMSON DERWENT on STN

ACCESSION NUMBER:

1997-535573 [49] WPIDS

CROSS REFERENCE:

2003-362085 [34]

DOC. NO. CPI:

C1997-171219

TITLE:

Treatment of bipolar disorder uses omega-3 fatty acids and phosphatidyl-choline - preferably eicosapentanoic acid or docosahexanoic acid or glycerol esterified with

omega-3 acids and phosphocholine.

DERWENT CLASS:

INVENTOR(S):

SEVERUS, W E; STOLL, A L

PATENT ASSIGNEE(S):

(BGHM) BRIGHAM & WOMENS HOSPITAL; (SEVE-I) SEVERUS W E;

(STOL-I) STOLL A L

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK	
WO 9739759 A2 19971030 (1997 RW: AT BE CH DE DK ES FI FR	49) * EN 13 A61K033-14 <
W: AU CA JP US AU 9727384 A 19971112 (1998	11) A61K033-14
US 6344482 B1 20020205 (2002	
US 2002091103 A1 20020711 (2002	48) A61K031-685

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9739759 AU 9727384 US 6344482 US 20020911	A2 A B1 03 A1 Cont of	WO 1997-US6712 AU 1997-27384 WO 1997-US6712 US 1999-269361 US 1999-269361	19970423 19970423 19970423 19990322 19990322
		US 2002-68035	20020205

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9727384	A Based on	WO 9739759
US 6344482	B1 Based on	WO 9739759

PRIORITY APPLN. INFO: US 1996-16140P

19960424; US 1999-269361

19990322

REFERENCE PATENTS:

No-SR. Pub

10/477,971 083-913

INT. PATENT CLASSIF.:

MAIN:

A61K031-20; A61K031-685; A61K033-14

SECONDARY:

A61K031-14; A61K031-203; A61K031-66; A61K033-00

BASIC ABSTRACT:

WO 9739759 A UPAB: 20030529

Treatment for bipolar disorder comprises administration of an Omega -3 fatty acid in sufficient dosage to reduce or eliminate symptoms. Also claimed are: (i) an Omega -3 phosphatidylcholine, used in the treatment of bipolar disorders, comprising of glycerol, where: (a) the alpha and beta carbons of the glycerol are both esterified to a fatty acid, at least one of which is an Omega -3 fatty acid; and (b) the gamma carbon of the glycerol is esterified to phosphocholine; and (b) a kit comprising a first component containing an Omega -3 fatty acid, and a second component containing an agent used in the treatment of bipolar disorders.

USE - The treatment is used to reduce or eliminate the symptoms of bipolar disorder (claimed). The Omega -3 fatty acids or Omega -3 phosphatidyl cholines are administered to bipolar patients in increasing doses until maximum benefit is achieved, preferably orally. Parenteral, transdermal, sublingual, or buccal administration is possible, or implantation.

ADVANTAGE - Omega -3 fatty acids inhibit thrombosis and platelet aggregation and can lower blood pressure. The Omega -3 phosphatidylcholines produce the same effect as lecithin due to the release of free choline, but reduce rather than increase risks of cardiovascular disease.

Dwg.0/0

FILE SEGMENT:

CPI

FIELD AVAILABILITY:

AB; DCN

MANUAL CODES:

CPI: B05-B01P; B10-C04E; B10-G02; B14-F02A; B14-F04

=> d his; d tot ibib abs clm

(FILE 'USPATFULL' ENTERED AT 06:18:21 ON 26 JAN 2004)

DEL HIS

L1 118576 S DEPRESSION

L2 1158 S (OMEGA(2W)(FATTY ACID#))

L3 7717 S PHOSPHATIDYLCHOLINE

18 S L1 AND L2 AND L3

L5 8 S (OMEGA(3W) PHOSPHATIDYLCHOLINE#)

L6 3 S L1 AND L5

L6 ANSWER 1 OF 3 USPATFULL on STN

ACCESSION NUMBER:

2003:17078 USPATFULL

TITLE:

T.4

Omega-3 fatty acids in the treatment of

depression

INVENTOR(S):

Stoll, Andrew, Lincoln, MA, UNITED STATES

KIND

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PATENT INFORMATION:	US 2003012827	A1	20030116	
APPLICATION INFO.:	US 2002-83913		20020227	
RELATED APPLN. INFO.:	Continuation-in-r	art of	Sar No	TTC 100

NUMBER

on 22 Mar 1999, GRANTED, Pat. No. US 6344482 A 371 of International Ser. No. WO 1997-US6712, filed on 23 Apr

DVLE

1997, PENDING Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Choate, Hall & Stewart, Exchange Place, 53 State

10/477,971 083,913

Street, Boston, MA, 02109

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

27

LINE COUNT:

617

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to a method of treating patients with major depression by administering omega-3 fatty acids. These may be administered in a substantially purified form, as part of a pharmaceutical composition, or as part of a larger molecule, e.g., a triacylglycerol, which releases free fatty acid after ingestion by a patient.

The present invention is also directed to triacylglycerols which are esterified at the gamma cardon of glycerol to phosphocholine and at either the alpha or beta carbon of glycerol to an omega-3 fatty acid. These "omega-3 phoshatidylcholines" are also used in the treatment of patients with major depression.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

CLM What is claimed is:

- 1. A method of treating a human patient for unipolar major depression, comprising administering an omega-3 fatty acid to said patient at a dosage sufficient to reduce or eliminate the symptoms of unipolar major depression.
- 2. The method of claim 1, wherein said omega-3 fatty acid is administered at a dose of between about 1 and about 30 grams per day.
- 3. The method of claim 1, wherein said omega-3 fatty acid is in a substantially pure form.
- 4. The method of claim 1, wherein said omega-3 fatty acid is eicosapentanoic acid.
- 5. The method of claim 4, wherein said eicosapentanoic acid is administered at a dose of between about 2 and about 10 grams per day.
- 6. The method of claim 1, wherein said omega-3 fatty acid is docosahexanoic acid.
- 7. The method of claim 6, wherein said docosahexanoic acid is administered at a dose of between about 1 and about 5 grams per day.
- 8. The method of claim 1, wherein said omega-3 fatty acid is alpha-linolenic acid.
- 9. The method of claim 1, further comprising administering a pharmaceutically effective dose of at least one member of lithium, a pharmaceutical antidepressant, an herbal antidepressant, an anticonvulsant, a mood stabilizer, an antipsychotic agent, and a benzodiazepine.
- 10. An omega-3 phosphatidylcholine useful in the treatment of unipolar major depression consisting of glycerol, wherein: a) the .alpha. and .beta. carbons of said glycerol are both esterified to a fatty acid, at least one of which is an omega-3 fatty acid; and b) the .gamma. carbon of said glycerol is esterified to phosphocholine.
- 11. The omega-3 phosphatidylcholine of claim 10, wherein both the .alpha. and .beta. carbons of said glycerol are

esterified to an omega-3 fatty acid.

- 12. The omega-3 phosphatidylcholine of either claim
 10 or 11, wherein eicosapentanoic acid is esterified to a member of the .alpha. carbon, the .beta. carbon, and both the .alpha. and .beta. carbons of said glycerol.
- 13. The omega-3 phosphatidylcholine of either claim 10 or 11, wherein docosahexanoic acid is esterified to a member of the .alpha. carbon, the .beta. carbon, and both the .alpha. and .beta. carbons of said glycerol.
- 14. The omega-3 phosphatidylcholine of either claim 10 or 11, wherein alpha-linolenic acid is esterified to a member of the .alpha. carbon, the .beta. carbon, and both the .alpha. and .beta. carbons of said glycerol.
- 15. The omega-3 phosphatidylcholine of claim 10, wherein eicosapentanoic acid is esterified to the .alpha. carbon of said glycerol and docosahexanoic acid is esterified to the .beta. carbon of said glycerol.
- 16. The omega-3 phosphatidylcholine of claim 10, wherein docosahexanoic acid is esterified to the .alpha.carbon of said glycerol and eicosapentanoic acid is esterified to the .beta. carbon of said omega-3 phosphatidylcholine.
- 17. A pharmaceutical composition comprising the omega-3 phosphatidylcholine of claim 10, wherein one or more unit doses of said composition provides an amount of said omega-3 phosphatidylcholine sufficient to reduce or eliminate the symptoms of unipolar major depression.
- 18. The pharmaceutical composition of claim 16, further comprising a member of lithium, a pharmaceutical antidepressant, an herbal antidepressant, an anticonvulsant, a mood stabilizer, an antipsychotic agent, and a benzodiazepine.
- 19. A method of treating unipolar major depression in a human patient, comprising administering the omega-3 phosphatidylcholine of claim 10 to said patient at a dose sufficient to reduce or eliminate the symptoms of unipolar major depression.
- 20. The method of claim 19, further comprising administering a pharmaceutically effective dose of at least one member of lithium, a pharmaceutical antidepressant, an herbal antidepressant, an anticonvulsant, a mood stabilizer, an antipsychotic agent, and a benzodiazepine.
- 21. A kit comprising a carrier containing in close confinement therein one or more components, wherein: a) a first component contains an omega-3 fatty acid; and b) a second component contains a psychotropic medication useful in the treatment of unipolar major depression
- 22. The kit of claim 21 wherein: a) said first component contains an omega-3 fatty acid selected from the group consisting of eicosapentanoic acid, docosahexanoic acid, and alpha-linolenic acid; and b) said second component is selected from the group consisting of lithium, pharmaceutical antidepressant, an herbal antidepressant, an

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> anticonvulsant, a mood stabilizer, an antipsychotic agent, and a benzodiazepine.

- 23. A kit comprising a carrier containing in close confinement therein, none or more components wherein: a) a first component contains an omega-3 phosphatidyl-choline; and b) a second component contains a psychotropic agent useful in the treatment of unipolar major depression.
- 24. The kit of claim 23, wherein the .alpha. carbon of said glycerol is esterified to eicosapentanoic acid and the .beta. carbon of said glycerol is a esterified to doocosa-hexanoic acid.
- 25. The kit of claim 23, wherein the .alpha. carbon of said glycerol is esterified to docosahexanoic acid and the .beta. carbon of said glycerol is a esterified to eicosapentanoic acid.
- 26. The kit of claim 23, wherein a member of eicosapentanoic acid, docosapentanoic acid, and alpha-linolenic acid is esterified to a member of the .alpha. carbon, the .beta. carbon, and both the .alpha. and .beta. carbons of said glycerol.
- 27. The kit of any one of claims 23-26, wherein said second component is selected from the group consisting of lithium, pharmaceutical antidepressant, an herbal antidepressant, an anticonvulsant, a mood stabilizer, an antipsychotic agent, and a benzodiazepine.

ANSWER 2 OF 3 USPATFULL on STN

ACCESSION NUMBER:

2002:172346 USPATFULL

TITLE:

Omega-3 fatty acids and omega-3

phosphatidylcholine in the treatment of bipolar

INVENTOR(S):

Stoll, Andrew L., Lincoln, MA, UNITED STATES

Severus, Wolfram E., Berlin, GERMANY, FEDERAL REPUBLIC

NUMBER KIND DATE US 2002091103 A1 20020711

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

US 2002-68035 A1 20020205 (10)

Continuation of Ser. No. US 1999-269361, filed on 22 Mar 1999, PATENTED

> NUMBER DATE WO 1997-US6712 19970423

PRIORITY INFORMATION:

Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Choate, Hall & Stewart, Exchange Place, 53 State

Street, Boston, MA, 02109

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

25 1

LINE COUNT:

439

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AΒ The present invention is directed to a method of treating patients with bipolar disorder by administering omega-3 fatty acids. These may be administered in a substantially purified form, as part of a pharmaceutical composition, or as part of a larger molecule, e.g. a triacylglycerol, which releases free fatty acid after ingestion by a patient.

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The present invention is also directed to triacylglycerols which are esterified at the gamma carbon of glycerol to phosphocholine and at either the alpha or beta carbon of glycerol to an omega-3 fatty acid. These "omega-3 phoshatidylcholines" are also used in the treatment of patients with bipolar disorder.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.
CLM What is claimed is:

- 1. A method of treating a human patient for bipolar disorder, comprising administering an omega-3 fatty acid to said patient at a dosage sufficient to reduce or eliminate the symptoms of said disorder.
- 2. The method of claim 1, wherein said omega-3 fatty acid is administered at a dose of between about 1 and about 30 grams per day.
- 3. The method of claim 1, wherein said omega-3 fatty acid is in a substantially pure form.
- 4. The method of claim 1, wherein said omega-3 fatty acid is eicosapentanoic acid.
- 5. The method of claim 4, wherein said eicosapentanoic acid is administered at a dose of between about 2 and about 10 grams per day.
- 6. The method of claim 1, wherein said omega-3 fatty acid is docosahexanoic acid.
- 7. The method of claim 6, wherein said docosahexanoic acid is administered at a dose of between about 1 and about 5 grams per day.
- 8. The method of claim 1, further comprising administering a source of lithium to said patient at a dose sufficient to reduce or eliminate the symptoms of said disorder.
- 9. The method of claim 1, further comprising administering a source of choline to said patient at a dose effective at reducing or eliminating the symptoms of said disorder.
- 10. An omega-3 phosphatidylcholine useful in the treatment of bipolar disorder consisting of glycerol, wherein: a) the alpha. and .beta. carbons of said glycerol are both esterified to a fatty acid, at least one of which is an omega-3 fatty acid; and b) the .gamma. carbon of said glycerol is esterified to phosphocholine.
- 11. The omega-3 phosphatidylcholine of claim 10, wherein both the .alpha. and .beta. carbons of said glycerol are esterified to an omega-3 fatty acid.
- 12. The omega-3 phosphatidylcholines of either claim 10 or 11, wherein eicosapentanoic acid is esterified to either the .alpha. or .beta. carbon of said glycerol.
- 13. The omega-3 phosphatidylcholine of either claim 10 or 11, wherein docosahexanoic acid is esterified to either the .alpha. or .beta. carbon of said glycerol.
- 14. The omega-3 phosphatidylcholine of claim 10, wherein eicosapentanoic acid is esterified to the .alpha. carbon of said glycerol and docosahexanoic acid is esterified to the .beta. carbon of said glycerol.

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- 15. The omega-3 phosphatidylcholine of claim 10, wherein docosahexanoic acid is esterified to the .alpha. carbon of said glycerol and eicosapentanoic acid is esterified to the .beta. carbon of said omega-3 phosphatidylcholines.
- 16. A pharmaceutical composition comprising the omega-3 phosphatidylcholine of claim 10, wherein one or more unit doses of said composition provides an amount of said omega-3 phosphatidylcholine sufficient to reduce or eliminate the symptoms of said bipolar disorder.
- 17. The pharmaceutical composition of claim 16, further comprising a source of lithium.
- 18. A method of treating bipolar disorder in a human patient, comprising administering the omega-3 phosphatidyicholine of claim 10 to said patient at a dose sufficient to reduce or eliminate the symptoms of said disorder.
- 19. The method of claim 18, further comprising administering a source of lithium to said patient at a dosage sufficient to reduce or eliminate the symptoms of said disorder.
- 20. A kit comprising a carrier containing enclosed confinement therein one or more components, wherein: a) a first component contains an omega-3 fatty acid; and b) a second component contains a therapeutic agent useful in the treatment of bipolar disorder.
- 21. The kit of claim 20 wherein: a) said first component contains an omega-3 fatty acid selected from the group consisting of eicosapentanoic acid and docosahexanoic acid; and b) said second component is selected from the group consisting of a source of choline and a source of lithium.
- 22. A kit comprising a carrier containing in close confinement therein, none or more components wherein: a) a first component contains an omega-3 phosphatidyl-choline; and b) a second component contains a therapeutic agent useful in the treatment of bipolar disorder.
- 23. The kit of claim 22, wherein the .alpha. carbon of said glycerol is esterified to eicosapentanoic acid and the .beta. carbon of said glycerol is a esterified to docosa-hexanoic acid.
- 24. The kit of claim 22, wherein the .alpha. carbon of said glycerol is esterified to docosahexanoic acid and the .beta. carbon of said glycerol is a esterified to eicosapentanoic acid
- 24. The kit of any one of claims 22-24, wherein said second component is selected from the group consisting of a source of choline and a source of lithium.

L6 ANSWER 3 OF 3 USPATFULL on STN

ACCESSION NUMBER: 2002:24309 USPATFULL

TITLE:

Omega-3 fatty acids in the treatment of bipolar

disorder

INVENTOR(S):

Stoll, Andrew L., 35 Old Winter St., Lincoln, MA,

United States 01773

Severus, Wolfram E., Badensche Strasse 7, D-10825

Berlin, GERMANY, FEDERAL REPUBLIC OF

NUMBER KIND DATE ______ ~----US 6344482 B1 20020205 PATENT INFORMATION: WO 9739759 19971030 19990322 US 1999-269361 APPLICATION INFO.: (9) WO 1997-US6712 19970423 19990322 PCT 371 date

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Jarvis, William R. A. LEGAL REPRESENTATIVE: Choate, Hall & Stewart

NUMBER OF CLAIMS: 9 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 387

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention is directed to a method of treating patients with bipolar disorder by administering omega-3 fatty acids.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

CLM What is claimed is:

- 1. A method of treating a human patient for bipolar disorder, comprising administering an omega-3 fatty acid to said patient at a dosage sufficient to reduce or eliminate the symptoms of said disorder, wherein said symptoms are symptoms of mania and symptoms of **depression**
- 2. The method of claim 1, wherein said omega-3 fatty acid is administered at a dose of between about 1 and about 30 grams per day.
- 3. The method of claim 1, wherein said omega-3 fatty acid is in a substantially pure form.
- 4. The method of claim 1, wherein said omega-3 fatty acid is eicosapentanoic acid.
- 5. The method of claim 4, wherein said eicosapentanoic acid is administered at a dose of between about 2 and about 10 grams per day.
- 6. The method of claim 1, wherein said omega-3 fatty acid is docosahexanoic acid.
- 7. The method of claim 6, wherein said docosahexanoic acid is administered at a dose of between about 1 and about 5 grams per day.
- 8. The method of claim 1, further comprising administering a source of lithium to said patient at a dose sufficient to reduce or eliminate the symptoms of said disorder.
- 9. The method of claim 1, further comprising administering a source of choline to said patient at a dose effective at reducing or eliminating the symptoms of said disorder.